Serial No.: To be assigned Case No.:

21020P

Page

310/500811

DT04 Rec'd PCT/PT0 0 7 JUL 2004

Amendments to the Claims:

This listing of claims will replace all prior versions, and listings, of claims in the application:

Listing of Claims:

A compound of Formula I 1. (original)

$$\begin{array}{c|c}
A & R^1 \\
 & R^2 \\
 & R^2
\end{array}$$

$$\begin{array}{c|c}
R^3 \\
 & R^4)_{0-4} \\
 & R^5 \\
 & R^5
\end{array}$$

or a pharmaceutically acceptable salt or hydrate thereof, wherein:

Ar is phenyl or naphthyl;

A is selected from: -CO₂H, 1*H*-tetrazol-5-yl, -PO₃H₂, -PO₂H₂, -SO₃H, and -PO(R⁵)OH, wherein R⁵ is selected from the group consisting of: C₁₋₄alkyl, hydroxyC₁₋ 4alkyl, phenyl, -C(O)-C₁-3alkoxy and -CH(OH)-phenyl, said phenyl and phenyl portion of -CH(OH)-phenyl optionally substituted with 1-3 substituents independently selected from the group consisting of: hydroxy, halo, -CO₂H, C₁-4alkyl, -S(O)_kC₁-3alkyl, wherein k is 0, 1 or 2, C₁₋₃alkoxy, C₃₋₆ cycloalkoxy, aryl and aralkoxy, the alkyl portions of said C₁₋₄alkyl, -S(O)_kC₁₋₃alkyl, C₁₋₃alkoxy and C₃₋₆ cycloalkoxy optionally substituted with 1-3 halo groups;

n is 2, 3 or 4;

each R¹ and R² is each independently selected from the group consisting of: hydrogen, halo, hydroxy, -CO₂H, C₁-6alkyl and phenyl, said C₁-6alkyl and phenyl optionally substituted with 1-3 halo groups;

R³ is selected from the group consisting of: hydrogen and C₁-4alkyl, optionally substituted with 1-3 hydroxy or halo groups;

Serial No.: To be assigned

Case No.:

21020P

Page

4

each R^4 is independently selected from the group consisting of: hydroxy, halo, -CO₂H, C₁₋₄alkyl, -S(O)_kC₁₋₃alkyl, wherein k is 0, 1 or 2, C₁₋₃alkoxy, C₃₋₆ cycloalkoxy, aryl and aralkoxy, the alkyl portions of said C₁₋₄alkyl, -S(O)_kC₁₋₃alkyl, C₁₋₃alkoxy and C₃₋₆ cycloalkoxy optionally substituted with 1-3 halo groups;

C is selected from the group consisting of:

- (1) C_{1-8} alkyl, C_{1-8} alkoxy, -(C=O)- C_{1-6} alkyl or -CHOH- C_{1-6} alkyl, said C_{1-8} alkyl, C_{1-8} alkoxy, -(C=O)- C_{1-6} alkyl and -CHOH- C_{1-6} alkyl optionally substituted with phenyl, and
- phenyl or HET, each optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, phenyl, C₁-4alkyl and C₁-4alkoxy, said C₁-4alkyl and C₁-4alkoxy groups optionally substituted from one up to the maximum number of substitutable positions with a substituent independently selected from halo and hydroxy, and said phenyl optionally substituted with 1 to 5 groups independently selected from the group consisting of: halo and C₁-4alkyl, optionally substituted with 1-3 halo groups,

or C is not present;

when C is not present then B is selected from the group consisting of: phenyl, C5-16alkyl, C5-16alkynyl, -CHOH-C4-15alkyl, -CHOH-C4-15alkynyl, C4-15alkynyl, -CHOH-C4-15alkynyl, C4-15alkoxy, -O-C4-15alkenyl, -O-C4-15alkynyl, C4-15alkylthio, -S-C4-15alkenyl, -S-C4-15alkynyl, -CH2-C3-14alkoxy, -CH2-O-C3-14alkenyl, -CH2-O-C3-14alkynyl, -(C=O)-C4-15alkyl, -(C=O)-C4-15alkynyl, -(C=O)-O-C3-14alkyl, -(C=O)-O-C3-14alkynyl, -(C=O)-N(R6)(R7)-C3-14alkyl, -(C=O)-N(R6)(R7)-C3-14alkyl, -(C=O)-C3-14alkyl, -N(R6)(R7)-C3-14alkynyl, -N(R6)(R7)-(C=O)-C3-14alkyl, -N(R6)(R7)-(C=O)-C3-14alkyl, -N(R6)(R7)-(C=O)-C3-14alkynyl, -N(R6)(R7)

when C is phenyl or HET then **B** is selected from the group consisting of: C_{1-6} alkyl, C_{1-5} alkoxy, -(C=O)- C_{1-5} alkyl, -(C=O)- C_{1-4} alkyl, -(C=O)- $N(R^6)(R^7)$ - C_{1-4} alkyl, C_{1-3} alkyl

Serial No.: To be assigned

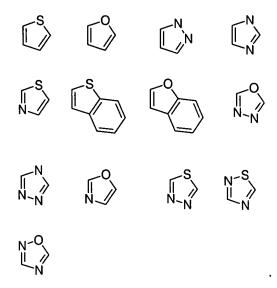
Case No.: 21020P

Page 5

when C is C_{1-8} alkyl, C_{1-8} alkoxy, -(C=O)- C_{1-6} alkyl or -CHOH- C_{1-6} alkyl then B is phenyl; and

 R^6 and R^7 are independently selected from the group consisting of: hydrogen, C_1 -galkyl and -(CH₂)_p-phenyl, wherein p is 1 to 5 and phenyl is optionally substituted with 1-3 substituents independently selected from the group consisting of: C_1 -galkyl and C_1 -galkoxy, each optionally substituted with 1-3 halo groups.

2. (original) The compound according to Claim 1 wherein HET is selected from the group consisting of:



- 3. (original) The compound according to Claim 1 wherein n is 2.
- 4. (original) The compound according to Claim 1 wherein n is 3.
- 5. (original) The compound according to Claim 3 wherein each R¹ and R² is independently selected from the group consisting of: hydrogen, -CO₂H, hydroxy, halo, C₁-3alkyl and phenyl.
 - 6. (original) The compound according to Claim 1 wherein A is PO₃H₂.
 - 7. (original) The compound according to Claim 1 wherein A is -CO₂H.

To be assigned Serial No.: 21020P

Case No.:

Page

The compound according to Claim 1 wherein A is $PO(R^5)OH$, 8. (original) wherein R5 is selected from the group consisting of: C1-4alkyl, hydroxyC1-4alkyl, C(O)-C1-2alkoxy and benzyl, wherein both the methyl and phenyl portions of said benzyl are optionally substituted with 1-3 halo or hydroxy groups.

- The compound according to Claim 1 wherein A is PO₂H₂. 9. (original)
- 10. (original) The compound according to Claim 1 wherein A is 1H-tetrazol-5-yl.
- 11. (original) The compound according to Claim 1 wherein R3 is hydrogen or methyl.
- 12. (original) The compound according to Claim 1 wherein each R4 is independently selected from the group consisting of: halo, hydroxy, C1-3alkyl, C1-3alkoxy, C1-3alkylthio, phenyl, benzyloxy and cyclopropyloxy.
- 13. (orignal) The compound according to Claim 1 wherein B is C₈₋₁₀alkyl and C is not present.
- 14. (original) The compound according to Claim 1 wherein B is C4-11alkoxy and C is not present.
- 15. (original) The compound of according to Claim 1 wherein **B** is phenyl, optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C₁-4alkyl and C₁-4alkoxy, and C is selected from the group consisting of: hydrogen, phenyl, C1-8alkyl, C1-8alkoxy, -(C=O)-C1-6alkyl and -CHOH-C₁₋₆alkyl, said C₁₋₈alkyl, C₁₋₈alkoxy, -(C=O)-C₁₋₆alkyl and -CHOH-C₁₋₆alkyl optionally substituted with phenyl.
- 16. (original) The compound according to Claim 1 wherein B is selected from the group consisting of: -CHOH-C₆-10alkyl, C₆₋₁₀alkylthio, -CH₂-C₅₋₉alkoxy, - $(C=O)-C_{6-10}alkyl,\ -(C=O)-O-C_{5-9}alkyl,\ -(C=O)-N(R^6)(R^7)-C_{5-9}alkyl,\ -N(R^6)(R^7)-(C=O)-N(R^6)(R^7)-(R^7)-(R^7)-(R^7)-(R^7)-(R^7)-(R^7)-(R^7)-(R^7)-(R^7)-(R^7)-(R^7)-(R^7)-(R^7)-(R^7)-(R^7)-(R^7)-(R^7)-(R^7)-(R^7)-(R^7$ C5-9alkyl, and C is not present.

Serial No.: To be assigned

Case No.: 21020P

Page

17. (original) The compound according to Claim 1 wherein $\bf B$ is C₁₋₆alkyl or C₁₋₅alkoxy and $\bf C$ is phenyl.

18. (original) The compound according to Claim 1 wherein B-C is

or

19. (original) The compound according to Claim 1 wherein Ar is phenyl and the group **-B-C** is attached to the phenyl ring at the 3- or 4-position.

20. (original) A compound of Formula Π

$$A \xrightarrow{\begin{array}{c} R^1 \\ C \\ R^2 \end{array} \xrightarrow{\begin{array}{c} R^3 \\ H \end{array}} (R^4)_{0-4}$$

$$B \\ C$$

$$II$$

or a pharmaceutically acceptable salt or hydrate thereof, wherein

the group **-B-C** is attached to the phenyl ring at the 3- or 4-position;

n is 2, 3 or 4;

Serial No.: To be assigned

Case No.: 21020P

Page

each R¹ and R² is independently selected from the group consisting of: hydrogen, -CO₂H, hydroxy, halo, C₁₋₃alkyl and phenyl, said C1-3alkyl and phenyl optionally substituted with 1-3 halo group;

A is selected from the group consisting of: 1H-tetrazol-5-yl, PO_2H_2 , PO_3H_2 , $-CO_2H$ and $PO(R^5)OH$, wherein R^5 is selected from the group consisting of: C_1 -4alkyl, hydroxy C_1 -4alkyl, C(O)- C_1 -2alkoxy and benzyl, wherein both the methyl and phenyl portions of said benzyl are optionally substituted with 1-3 halo or hydroxy groups;

R³ is hydrogen or methyl;

each R⁴ is independently selected from the group consisting of: halo, hydroxy, C₁₋₃alkyl, C₁₋₃alkoxy, C₁₋₃alkylthio, phenyl, benzyloxy and cyclopropyloxy; and

B-C is selected from the group consisting of:

- (1) **B** is C₈₋₁₀alkyl and **C** is not present.
- (2) \mathbf{B} is C₄₋₁₁alkoxy and \mathbf{C} is not present.
- (3) **B** is phenyl, optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C_{1-4} alkyl and C_{1-4} alkoxy, and **C** is selected from the group consisting of: hydrogen, phenyl, C_{1-8} alkyl, C_{1-8} alkoxy, -(C=O)- C_{1-6} alkyl and -CHOH- C_{1-6} alkyl, said C_{1-8} alkyl, C_{1-8} alkoxy, -(C=O)- C_{1-6} alkyl and -CHOH- C_{1-6} alkyl optionally substituted with phenyl;
- (4) **B** is -CHOH-C₆-10alkyl, C₆₋₁₀alkylthio, -CH₂-C₅-9alkoxy, -(C=O)-C₆₋₁₀alkyl, -(C=O)-O-C₅-9alkyl, -(C=O)-N(R⁶)(R⁷)-C₅-9alkyl or -N(R⁶)(R⁷)-(C=O)-C₅-9alkyl, and **C** is not present.
 - (5) **B** is C_{1} -6alkyl or C_{1} -5alkoxy and **C** is phenyl.
 - (6) **B-C** is

Serial No.: Case No.: Page

9

or

21. (original) A compound selected from the group consisting of:

OH HO—==0
CH ₃
CH ₃ OH OH
CH ₃ OH
HO OH
CH ₃
CH ₃ OH OH
CH ₃ OH
CH ₃ OH
CH ₃ OH

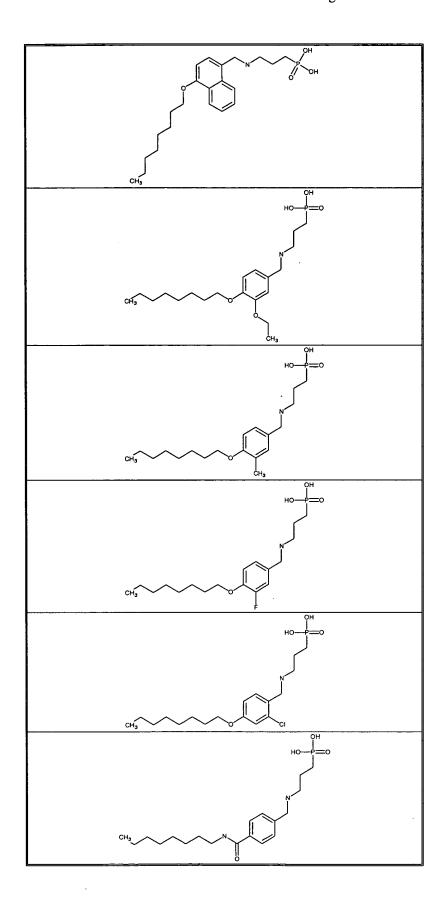
Serial No.: Case No.:

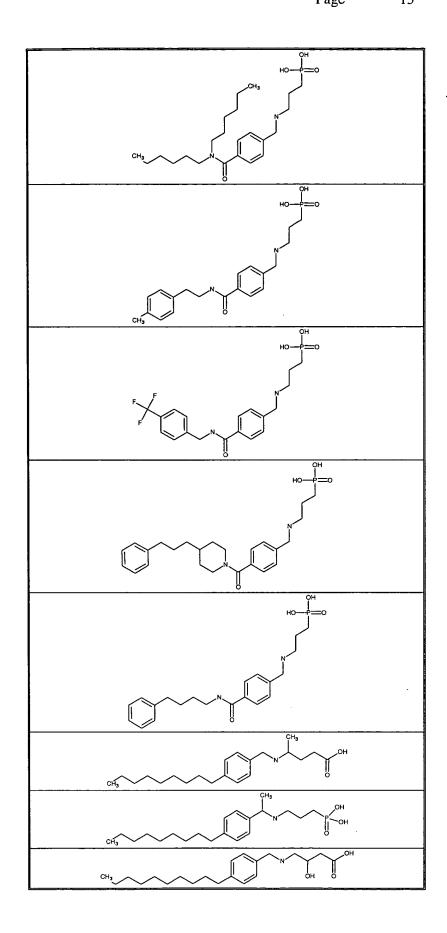
Page 11

QH (Non)
OH OH
сң,
CH ₂ S
N N N
CH ₃ V V V
↑ ↑ ↑ NH
CH ₃
N OH
CH, V V V
CH ₃ F B
OH I
но—ф==0
N ^r
CH ₃
CH ₃
ОН
но
_
CH ₉ N
CH ₃ CH ₃
CH ₃ CH ₃ OH
но
1
Br
CH ₉
Br OH
но——
l N
CH ₃
V V

Serial No.: Case No.: Page

12





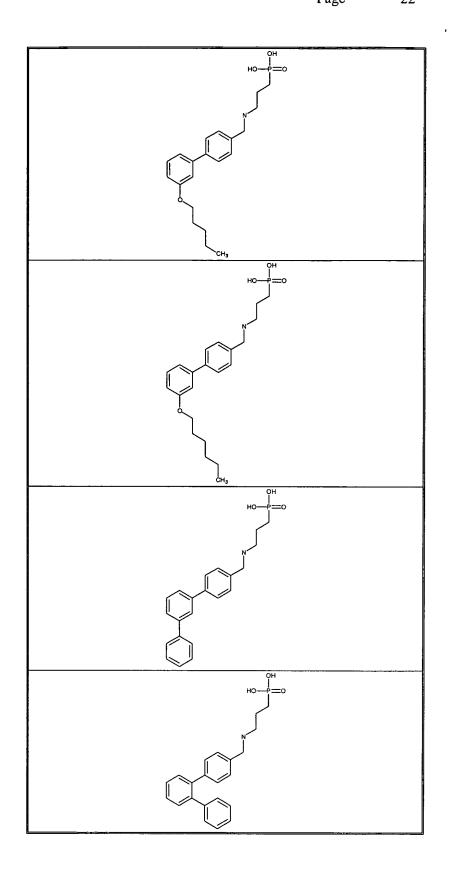
Serial No.: Case No.: Page To be assigned 21020P

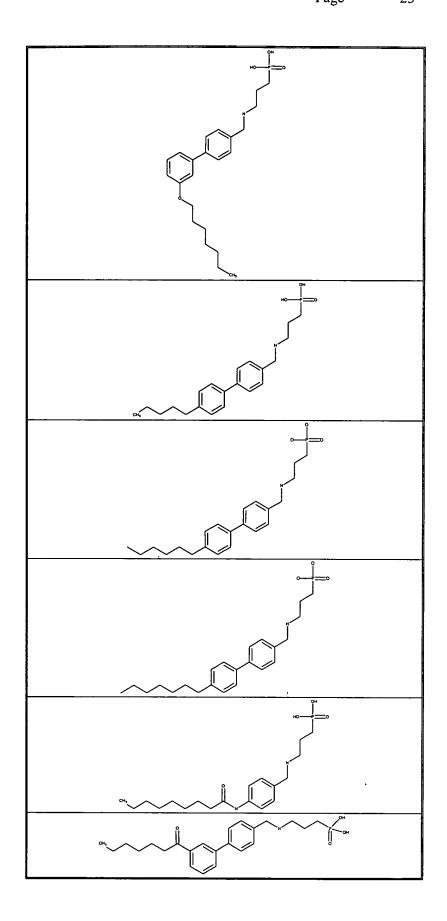
CH ₃ OH
CH ₃ OH OH
CH ₃ OH
CH ₃ OH
CH ₃ OH
CH ₃
CH ₃ OH
CH ₃
CH ₃ OH OH
CH ₃ OH
CH ₃ OH
CH ₃
CH ₃ OH
CH ₃ OH

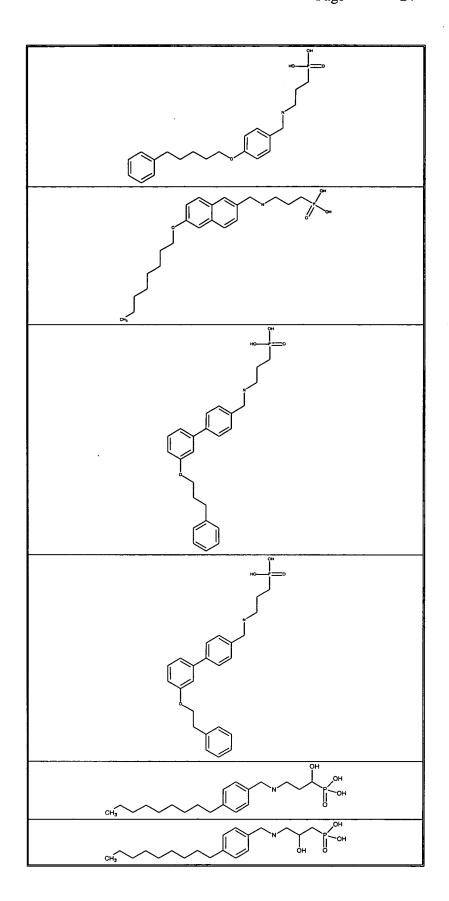
CH3 OH OH
CH ₃
но
CH ₉
OH HO—=O
CH ₉
НО
CH ₉
но
CH ₉
но
CH ₃

HO—P=O
OH HO—==0
CH ₃
OH HO—==0
CH ₃ OH HO PO
CH ₃ OH ₃ OH
HO—J=O
CH ₃

CH ₃ CH ₃ CH ₃ CH ₃
OH HO—P=O
CH ₃ CH ₃ OH HO
CH ₃ CH ₃
HO PHO
CH ₃







М № ОН Н ОН
_F
О СН,
N OH
a H
CH ₃
N B OH
О СН3
N OH
OH ₃
Н ОН
s
CH ₃ CH ₃
CH ₃ CH ₃
OH OH
CH ₃
CH3 OH
N C H _d
CH ₃ OH
CH ₃ OH

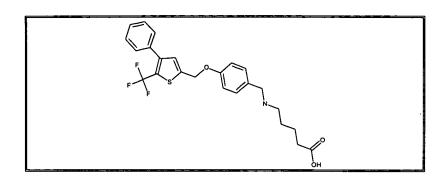
Serial No.: Case No.: To be assigned 21020P

Page 27

F S N N OH OH
F P OH
F F F
CH ₉ CH ₉ OH
F S N OH
F S OH
F S N OH OH CH ₃
F S OH OH

Serial No.: To be assigned

Case No.: 21020P Page 29



22. (currently amended) A method of treating oan an immunoregulatory abnormality in a mammalian patient in need of such treatment comprising administering to said patient a compound in accordance with Claim 1 in an amount that is effective for treating said immunoregulatory abnormality.

23 to 33 (canceled).

- 34. (original) A method of suppressing the immune system in a mammalian patient in need of immunosuppression comprising administering to said patient an immunosuppressing effective amount of a compound of Claim 1.
- 35. (original) A pharmaceutical composition comprised of a compound in accordance with Claim 1 in combination with a pharmaceutically acceptable carrier.